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Honey in Medicine

Stefan Bogdanov

SHORT HISTORY OF HONEY IN MEDICINE



Pabasa tombs, 26th Dynasty, 760-656 BC





Preparation of honey medicine from Materia Medica, Dioscorides, Arab translation 1224

Wound healing was probably the first use of honey for human health. In the oldest human scriptures from Sumer, dating back about 2000 BC a prescription for treating wounds states: *Grind to a powder river dust and* (words missing) then knead it in water and honey and let plain oil and hot cedar oil be spread over it⁷⁴

According to the Ebers papyrus (1550 BC) honey is included in 147 prescriptions in external applications: "*Mix honey, red ochre, powdered alabaster to cure spotted baldness*" or includes honey after surgery, as suppository and to reduce inflammation. ⁷⁴

According to the Smith papyrus (1700 BC) it was used in wound healing: "Thou shouldst bind [the wound] with fresh meat the first day [and] treat afterwards with grease, honey [and] lint every day until he recovers." ⁷⁴

In the first compendium of ancient Chinese Medicine Shen Nang, compiled many years BC, and mentioned in a written form for the first time around 200 AD there are many prescriptions and medical indications which contain honey¹³⁹.

In ancient India ayuruvedic medicine uses honey for many purposes. According to the Ayruveda classic Ashtanga Hridaya, written about 500 AD honey can be used against many diseases, e.g. healing and cleaning wounds, against different internal and external infections⁸¹

The ancient Greeks considered honey as medicine and believed that if bee honey is taken regularly human life could be prolonged. Early thinkers such as Homer, Pythagoras, Ovid, Democritus, Hippocrates and Aristotle mentioned that people should eat honey to preserve their health and vigour. Dioscorides, in the first century AD (see picture to the left) used honey for treating wounds⁸⁴

Honey was the most useful substance used in old Roman pharmacopoeia. Pliny writes that it is good for afflictions of the mouth, pneumonia, pleurisy and snake bites⁷⁴

The wise Solomon praises the virtues of honey in the old testament. The Koran says" thy Lord taught the bee to build its cells in hills, on trees and in (men's) habitations..... there issues from within their bodies a drink of varying colours, wherein is healing for mankind (Quran 16:68-69).

The ancient Maya civilisations used Melipona (stingless bee) honey in the treatment of cataracts ⁷⁴

Today the knowledge on the healing virtues of honey and the other bee product is called apitherapy is compiled in many books or on the Internet www.apitherapy.com, www.apitherapy.org

HONEY IN TRADITIONAL MEDICINE

Honey in traditional Chinese medicine

Honey was mentioned as medicine by Shen Nang, some 2000 BC. In the "52 prescription book,. 3th century BC includes a prescription including honey. According to Chinese medicine honey has a balanced character (neither Yin nor Yang) and acts according to the principles of the Earth element, entering the lung, spleen and large intestine channels. There are many original prescriptions and medical indications which contain honey¹³⁹

Christopher Gussa a TCM practioneer writes on www.naturalnews.com :

"Honey is known as Feng Mi and has the ability to nourish yin energy and strengthen the spleen. Apart from its widely recognized nutritional value, honey is also the Chinese people's favorite as a "neutral" food with medicinal properties. In the "Compendium of Materia Medica," the TCM classic by pharmacist Li Shizhen in the Ming Dynasty (1368-1644), "Honey can help dispel pathogenic heat, clear away toxins, relieve pain and combat dehydration." Li Shizhen showed that eating honey regularly resulted in clear sight and rosy cheeks. He also wrote that eating honey every morning can help prevent constipation and it is a good choice for those who suffer chronic coughing.

TCM also shows that due to honey's affinity for the stomach and spleen it can greatly enhance the effect of many of the super-tonics such as He Shou Wu"

According to the Fundamentals of Chinese Medicine honey has following properties

• "balanced, sweet, non-toxic. Enters the lung, spleen and the large intestine meridian channels. Supllements the center and moistens the lung. Relieves pain and resolves toxin.....Treats cough due to lung dryness; constipation due to dryness of the intestines; stomach pain; deep source nasal congestion, mouth sores, scalds and burns" ²²

Sui Wan summirises: "Honey has been used in traditional Chinese Medicine to treat many diseases for more than two thousands years. In Traditional Chinese Medication, honey is a combination used with bee venom, propolis, royal jelly, pollen and other herbal medications in medical treatment. In addition, the following diseases are good indications for using honey. I) Infectious Diseases: Bacterial Infections: A. Infections caused by gram-positive bacteria, e.g. Streptococcal infections- Pharyngitis, Enterococcal infections, Pneumococcal infections; Staphyloccus Aureus infections, B. Infections caused by gram-negative bacteria, e.g. Meningococcal infections- Meningococcal meningitis., Salmonellae infections-Typhoid fever, Salmonella Gastro-enteritis., Shigella infections-Shigella dysentery. II) Gastrointestinal Diseases: A. Gastritis, B. Peptic ulcer Disease, C. Celiac Disease, D. Antibiotic-Associated Colitis. E. Inflammatory Bowel Diseases. III) Allergic and Immunologic Disorders: A. Allergic Rhinitis B. Rheumatoid arthritis, C. Systemic Lupus Erythematous(SLE), D. Ankylosing spondylitis, E. Multiple Sclerosis. IV) Traumatic wounds: traumatic and surgical wounds managements" ¹⁴³

Honey in Ayurveda Medicine

Honey has a long tradition in traditional ayurveda medicine. This topic has been reviewed by 126

According to D Ramanathan, director of the Sitaram Ayurveda Pharmacy Limited & Specialty Hospital, Thrissur on the role honey plays in ayurvedic treatment: "Honey known as madhu in ayurvedic scriptures is one of the most important medicines used in ayurveda. In ayurveda, honey is used for both internal and external applications. It is mainly used for the treatment of eye diseases, cough, thirst, phlegm, hiccups, blood in vomit, leprosy, diabetes, obesity, worm infestation, vomiting, asthma, diarrhoea and healing wounds. It is also used as a natural preservative and sweetener in many ayurvedic preparations. It is also used as a vehicle along with some medicines to improve its efficacy or to mitigate the side effects of the other medicines it is mixed with. It is also known to mitigate the increased kapha dosha. (Kapha dosha is the ayurvedic category for body constitutions- those with kapha dosha are of larger proportions with robust frame.) It should also be kept in mind that fresh honey helps to increase body mass while old honey produces constipation and decreases body mass. Honey should not be heated or consumed warm as it causes toxic effect. Cold honey should always be preferred.

According to ayurveda, there are eight different types of honey:

- 1.Makshikam: Used in the treatment of eye diseases, hepatitis, piles, asthma, cough and tuberculosis
- 2.Bhraamaram: Used in the treatment when blood is vomited

- 3.Kshoudram: Used in the treatment of diabetes
- 4. Pauthikam: Used in the treatment of diabetes and urinary infection
- 5. Chathram: Used in the treatment of worm infestation, when blood is vomited and diabetes
- 6. Aardhyam: Effective for eye diseases, cough and anaemia
- 7. Ouddalakam: Increases taste and swarasudhi. Used in the treatment of leprosy and poisoning cases
- 8.Daalam: It increases digestion and helps in the treatment of cough, vomiting and diabetes.

(The gathering of these honeys in described in ¹²⁶)

We prescribe a particular brand called Samskritha Madhu (which means cultured or purified honey) which is made by most of the authentic ayurvedic Manufacturing units as per the ayurvedic scriptures. In ayurveda, what is the bee species that is most preferred- are they are Italian species (that are kept in boxes) or Indian species (that are found in the wild). As far as Ayurveda is concerned our acharyas who made this system of medicine utilized Indian honey and tested the wild honey, hence any ayurvedic physician will prefer the Indian honey wild honey"

Honey in folk and traditional medicine

A traditional medicine branch, called apitherapy, has developed in recent years, offering treatments based on honey and the other bee products against many diseases. The knowledge on this subject is compiled in various books e.g. ^{34, 79, 124} or on relevant web pages such as www.apitherapy.com, www.apitherapy.org

Unifloral honey in practical apitherapy after³³

Honey type	Applications
Acacia, liquid and mild;	Sweetener for people with Diabetis Type II. Improved digestions. Applied at
Buckwheat, dark and strong	diseases of stomach, intestines, liver and kidney High antioxidant activity, improves digestions, to be taken by pregnant women
Buckwitch, dark and strong	and when nursing
Eucalyptus, dark and strong	Against infections and diseases of respiratory organs and urinary passages.
Callying doubt and strong	Increases immunity Lich entire ident activity. Invigorating at facigue and convelegence, against
Calluna, dark and strong	High antioxidant activity. Invigorating at fagigue and convalesence; against problems with kidney urinary bladder
Chestnut, dark and strong	Improves blood circulation; against anemia and infactions of kidney urinary
	bladder
Clover, light und mild	sedative
Lavendel, aromatic	Treatment of wounds, burns, insect stings, infections or respiritory organs and depressions
Linden, strong, aromatic	Diaphoretic, diuretic, palliative, apetising; against cold, flu, cough, sinusitis, headache, sleeplessness and anxiety.
Manuka, Dark and strong	High antibacterial activity, against infections and for wound healing
Dandelion, aromatic	Hemo-protective, against gastric, intestine, liver, kidney and gall bladder diseases
Citrus, light and mild	Against indigestion and sleeplessness
Rape, mild	Sedative, relaxing
Rosmarine, mild	Hemo-protective; against gastric, intestine, liver diseases
Sunflower, mild	spasmolytic in asthma cases, gastric, intestine colic
Fir, honeydew, dark and strong	High antioxidant activity. Against infections of respiritory organs
Thyme. dark and strong	Against infections of respiratory organs; wound treatment

Unifloral honeys are used in folk medicine for different purposes. The applications given in the table below remain to be confirmed by experimental science. Indeed, in most scientifically conducted clinical studies the botanical origin of the honey was not determined. On the other hand, the antibacterial and the antioxidant activity of honey depends strongly on the botanical origin. Health enhancing effects of different unifloral honey have been claimed in different practical apitherapy books, e.g. ^{32, 89, 100}. The table below has been compiled from them. At present there is no scientific explanation of many of the claimed effects.

HONEY AND WOUND CARE: AN OVERVIEW

By Keith Cutting, with some additions by S. Bogdanov

After Ostomy Wound Management⁴⁹,

History

Until the first part of the 20th century, honey dressings were part of everyday wound care practice. With the advent of antibiotics in the 1930s and 1940s, views changed and honey was consigned to items of historical interest. Misuse of antibiotics, the emergence of resistant bacteria, and increasing interest in therapeutic honey have provided an opportunity for honey to be re-established as a broad-spectrum, antibacterial agent that is non-toxic to human tissue.

Despite lack of promotional support from large corporations, interest in the use of honey in wound management has increased in recent years. However, a clinical profile in wound care commensurate with other modalities has not been achieved despite offering similar indications of use and an increase in research activity and clinical reports. It is observed that "The therapeutic potential of uncontaminated, pure honey is grossly underutilized" ¹⁶²

Clinicians need reassurance that any health-related agent is safe and meets its stated therapeutic purpose. Therefore, it is important to emphasize that although natural in origin, the honey used in wound care should be of medical-grade standard and not sourced from honey destined for the supermarket shelf. Medical grade honey is filtered, gamma-irradiated, and produced under carefully controlled standards of hygiene to ensure that a standardized honey is produced 157

Therapeutic benefit of honey in wound care

The therapeutic properties of honey are variable and depend on the type of honey used ⁹⁸. Manuka (the Maori name for the New Zealand tea tree/bush Leptospermum scoparium) or Leptospermum is honey derived from the tea tree; the former is the more widely used term. In a review of the literature, Moore showed that Manuka honey has "very special healing properties" and described it as "the best natural antibiotic in the world" ¹⁰⁵.

Medical grade Manuka honey is prepared purely for medical use and controlled by a rigorous set of systems and standards. These exacting standards apply to the leptospermum honey distributed in US (MedihoneyTM, Derma Sciences, Princeton, NJ). This product is a blend of *L. scoparium* (Manuka) and *L. polygalifolium know* as Jelly Bush.

A systematic review ¹⁰⁵ of honey as a wound dressing noting the dearth of good evidence on topical wound agents contradicts Molan's literature review ¹⁰⁰ of the evidence (17 randomized, controlled trials involving 1,965 participants and five clinical trials involving 97 participants, plus numerous case studies) — supports the use of honey as a wound dressing and underscores clinician failure to recognize that evidence. Molan's research reviews also addressed the range of honey's therapeutic activities ¹⁰²:

Bioactivity of honey	Suggested Rationale
Prevention of cross-contamination	Viscosity of honey provides a protective barrier
Provides a moist wound healing environment	Osmolarity draws fluid from underlying tissues
Dressings do not adhere to wound surface. Tissue does not grow into dressings	The viscous nature of honey provides an interface between wound bed and dressing
Promotes drainage from wound	Osmotic outflow sluices the wound bed
Removes malodor	Bacterial preference for sugar instead of protein (amino acids) means lactic acid is produced in place of malodorous compounds
Promotes autolytic debridement	Facilitates the autolytic action of proteases
Stimulates healing	Stimulation of cytokine production ¹³⁰
Anti-inflammatory	Number of inflammatory cells reduced in honey-treated wounds Prevention of inflammation by honey flavonoids ⁹⁰
Managing infection	Antiseptic properties found to be effective against a range of microbes including multi-resistant strains
Increases immune reaction	Effect of lipopolysacharides and apalbumine 1 and 2 ⁶³
Promotes pain relief and epitelisation	Analgetic effect of honey 87
Regulates oxidative stress in the wound	Inhibition of reactive oxygen species production by activating polymorphonuclear neutrophils ¹⁵³
Inhibits adhesion of bacteria in wound	Antiadhesive properties against wound pathogens ⁸⁹

Wound bed preparation

Wound bed preparation may be viewed as management of the wound in order to accelerate endogenous healing ¹⁰². The bio-activity of honey aligns closely with the concept of wound bed preparation. The physiology of healing in acute wounds is a carefully controlled series of events that ensures healing progresses in a timely fashion. However, in chronic wounds this orderly sequence is disrupted and the repair process is delayed. If wound bed preparation is to be successful, the impediments to healing must be recognized and addressed, implying appropriate management of exudate, devitalized tissue, and associated bioburden. The appropriate application of honey dressings offers a way forward in managing potential wound-related barriers to healing.

Exudate

A clinical study by Al-Waili and Saloom¹⁰ compared honey with topical antiseptics in 50 patients with postoperative abdominal wound infections; Ahmed et al's ⁶ non-randomizes study of 60 patients with chronic surgical or trauma wounds; and Betts and Molan's ³⁰ in vivo pilot study reported that honey helps reduce the amount of wound exudate. This is most likely a consequence of honey's anti-inflammatory properties. Inflammation — increased vessel permeability increases fluid movement into soft tissue, subsequently increasing surface exudate. A decrease in inflammatory cells has been found (histologically) in animal models following application of honey in full-thickness burns. Similar findings¹²³ have been reported in animal studies comparing ampicillin and nitrofuazone in treating infection of full-thickness wounds^{82,115}. The anti-inflammatory activity of honey also has been documented in clinical studies of human burn wounds and in "in vitro studies¹⁴⁸⁻¹⁵⁰. The potential consequences of effectively managing inflammation include rapid reduction of pain, edema, and exudate; additionally, hypertrophic scarring is minimized by avoiding protracted inflammation that may result in fibrosis¹⁰³. It follows that reducing inflammation lessens exudate production and dressing change frequency, which may conserve resources in terms of dressings used, staff time, and unnecessary disturbance of the patient and the wound bed.

Devitalized tissue

It has been established that dressings that create the type of moist wound environment that honey provides facilitate the process of autolytic debridement. The osmotic pull of honey draws lymph from the deeper tissues and constantly bathes the wound bed. Lymph fluid contains proteases that contribute to the debriding activity of honey. In addition, the constant sluicing of the wound bed is believed to help remove foreign body (e g, dirt and grit) contamination 103 . Molan 99 has suggested the most likely explanation for honey's debriding activity involves the conversion of inactive plasminogen to plasmin, an enzyme that breaks down the fibrin that tethers slough and eschar to the wound bed. Stephen-Haynes 147 who presented the results of three patient case studies and an additional five patients who benefited from management of wound malodor, attests to the clinical impact of honey in debridement. Malodor is known to occur in a variety of wounds in conjunction with slough and necrotic tissue; it is a particular concern when managing fungating lesions. Malodorous substances such as ammonia and sulphur compounds are produced when bacteria metabolize protein. Because honey provides bacteria an alternative source of energy (glucose), these noxious compounds are no longer produced and wound malodor is avoided.

Maceration

Macerated periwound skin can be a problem in some wounds and is often related to the dressing used⁴⁸. The osmotic action of honey, previously mentioned, has been shown in previous reviews of the literature to reduce the risk of maceration — honey draws moisture rather than donates it ¹⁰³. Thus, periwound skin is protected from overhydration.

Bioburden

Honey has been shown in clinical observations to have the ability to manage wound infection in situations where conventional antimicrobial (antibiotics/antiseptics) have failed^{55, 57, 158}. Honey also has been found to be effective in vitro against a range of multiresistant organisms including methicillin-resistant *Staphylococcus aureus* (MRSA), vancomycin-resistant Enterococci (VRE), and other multiresistant Gramnegative organisms including *Pseudomonas aeruginosa* ⁶⁴. Other in vitro studies involving different microorganisms also have demonstrated honey's effectiveness against antibiotic-resistant bacteria ^{46, 78}. George and Cutting specifically identified honey's antibacterial activity⁶⁴. The binding of water can be added to these antibacterial properties:

- The high sugar content/low water activity provides osmotic action
- Acidic pH (3.2 to 4.5) inhibits bacterial growth
- Glucose oxidase enzyme helps produce hydrogen peroxide
- Plant-derived factors (present in some honeys and not specifically identified).

The antibacterial action most likely reaches below the wound surface. Although topical honey manages superficial bacteria (bactericidal in action rather than bacteriostatic) ^{46, 154} it also has been shown in vitro to provide prompt clearance of deep-seated infection and boils with unbroken skin, suggesting that honey's antibacterial activity may diffuse through the skin to deeper tissues. Cooper et al ⁴⁶ performed sensitivity testing of 17 strains of *P. aeruginosa* isolated from infected burns using two honeys with different types of antibacterial activity; Wahdan¹⁵⁴ compared the antibacterial activity of a sugar solution and honey on 21 types of bacteria and two types of fungi.

Some commercial honey preparations used in wound healing



Medihoney for wound care



Antibacterial wound gel



Moistering cream agains eczemas



Medihoney wound gauze



Wound-healing creams



Wound dressing with Medihoney gauze

Biofilms

In recent years, attention has turned to the potential role of biofilms in wound infection. A biofilm may be described as a bacterial community living within a self- produced extracellular polysaccharide (EPS) matrix. The EPS protects the bacterial community from antimicrobial and phagocytic onslaught. Lately, in vitro evidence has indicated that honey is an effective agent for preventing biofilm formation. In an in vitro study it was found that laboratory-grown Pseudomonal biofilms were disrupted following application of Manuka honey⁷¹. These findings are particularly encouraging when considering the emergence of antimicrobial-resistant bacteria. No evidence has yet been presented that bacterial resistance to honey has occurred — it is highly unlikely that bacteria will select for resistance to honey because bacteria rely on sugar as a source of food.

Cross-contamination

Use of honey dressings may help prevent cross contamination. This is and will remain an important issue in healthcare. The viscous nature of honey is believed to provide a physical barrier that helps safeguard patients by preventing cross contamination.

Dressing wounds with honey

All dressings must be used in accordance with the manufacturer's instructions. This helps endorse the maxim do no harm and ensure that the full benefit of the product is realized.

Because of its fluid and viscous nature, honey can be difficult to apply. This is particularly true when profuse exudate is present, diluting the honey. Experience has shown that use of the appropriate honey vehicle, including a secondary dressing, can sometimes circumvent this problem:

- **Honey liquid or gel dispensed from a tube:** Applied directly onto wound or onto appropriate dressing before application
- **Absorbent alginate dressing with honey:** Can be applied to most acute/chronic wounds including infected or sloughy wounds
- **Honey in a hydrocolloid-like sheet:** Should be selected according to the exudate level of the wound **Alleroy**

Before honey is applied to a wound, the patient should be asked routinely if he/she is allergic to honey or bee products, including bee stings. It is advisable not to proceed with a honey-containing dressing if the answer is affirmative.

Discomfort

Occasionally, some patients report transient stinging on application of honey. The discomfort often disappears in a short period of time or after the first few applications. Analgesia is required only in those rare circumstances when pain may persist. In a review paper, Molan5 noted that patient response to honey applied to open wounds was reported as soothing, pain-relieving, and non-irritating, and demonstrated no adverse effects¹⁰⁰

Conclusion

The resurgence of interest in honey as a modern wound dressing offers opportunities for both patients and clinicians. Recent additions to the honey product range of dressings indicate commercial confidence in the future of therapeutic honey. The wheel has turned full circle and honey is being re-established as a valuable agent in modern wound care management. Its advantages — providing a moist environment, debriding, deodorizing, antibacterial, anti-inflammatory capabilities — are factors that have been shown to facilitate healing. These advantages have been experienced by patients and clinicians in Europe and Australia and are now available to patients in North America. Use of any medical device must be based on clinical justification and available evidence about product safety and effectiveness. Continued research is needed to increase our understanding about the role of honey in a variety of wounds and its effect on healing compared to other treatment modalities.

HONEY FOR WOUND HEALING UNDER HOME CONDITIONS

Besides scientifically based use of honey in wound care in hospitals (see next section), honey can also be used under home conditions, as it was used for many centuries. Although sterilised honey is only used in hospitals, raw honey can also be used under home conditions without any risk, as no adverse effects have been reported. Indeed, Prof. Descotte lectured in several Apitherapy conferences that his team has used raw honey routinely for wound care in hundreds of cases in the hospital of Limoge, France⁵³

Honey applied in wound healing in a Swiss hospital



Wound at the treatment beginning

A painful and infected wound on the left leg. The fracture was stabilised with plates and screws. After several operations the blood circulation of the leg was diminished and sores were built because of the prolonged bed lying.

The wound treatment with conventional means was not successful. The patient agreed to make a honey treatment.



After only 5 days of treatment the wound condition was significantly improved. New tissues were built and the bacterial inflammation has diminished significantly

After two months the wound was completely closed. The cicatrise is almost invisible and he skin is healthy, tender and elastic at the same time.

photos and comment by Kathrin Rieder, Switzerland, application see below

.Application of honey for wound healing under home conditions, after³³

It is not necessary to disinfect wound because honey will disinfect it

- 1. Apply honey as much honey on a gauze or clean cotton cloth as it is necessary to cover wound fully. Gauze and cloth need not be sterile.
- 2. Abscesses, cavity or deep wounds need more honey to adequately penetrate deep into the wound tissues. The wound bed should be filled with honey before applying the honey dressing pad.
- 3. Change bandage once a day. When doing it, wound need not be cleaned from honey. Honey is "dissolved" in the wound or sticks to the gauze.
- 4. When changing the bandage remove hornification at the border of the wound with a pincette. This can be done under running water or with a soft tooth brush. Cell debris, which were not removed will not disturb healing process.
- 5. After cleaning, wound should be padded with as much gauze as is needed for drawing the wound liquid.
- 6. If wound is infected by yeast or it heals badly, a mixture of honey-betadine 1:1 can be used.

The applications of honey in wound and burn healing have been treated in different reviews, the recent ones being: 15, 35, 36, 51, 75, 76, 86, 132

Honey against infections of the skin

Honey inhibits a number of microbes relevant in skin infections such as *Staph. aureus*, *Pseudomonas aur*, *E. coli*, *Candida alb*. and a number of dermatophytes and has thus a potential for application against microbisl skin diseases⁹⁴

Honey against eye diseases

Since ancient times honey has been used for the treatment of eye disorders. This topic is reviewed by Molan, 2001, see there the original references¹⁰²: Aristotle has written in his Historia Animalium that "honey is good as a salve for sore eyes". It has also been used by traditional Indian medicine and in Mali. In the Rangarya Medical College of India it has been used to treat corneal eye ulcers, treatments of plepharitis (inflammation of he eye-lids) catarrhal conjunctivitis and keratitits. Honey is also successful in various ailments of the cornea. The use of honey in Russia has been reviewed: undiluted or 20-50 % water solutions being applied to the eye under the lower eye lid against chemical and thermal burns of the eye, conjunctivitis and infections of the cornea. The healing effect of honey is explained by its anti-inflammatory, antibacterial and antifungal actions of honey.

There are reports on the successful treatment by honey of keratitis, conjunctivitis and blepharitis in Egypt⁵⁹.

A successful double blind trial on the efficacy of treatment of vernal keratoconjunctivitis has been reported ¹³⁴

The positive effect in keratitis to reducing the levels of angiogenic factors (VEGF and TGF-beta), inflammatory cytokines (IL-12) and chemokines (CC chemokine receptor 5(CCR-5)¹⁵²

Another explanation of the healing effect of honey in eye diseases is a irritation effect, triggering healing processes of the eye¹⁹. Stingless bee honey has been traditionally used by the Mayas against cataract¹⁰⁸.

ORAL HEALTH



There is much debate whether honey is harmful to teeth. Some reports show a cariogenic effect of honey^{37, 137}, while others claim that the effect of honey is less cariogenic effect that sucrose^{52 38}. Due to its antibacterial activity honey ingestion inhibits the growth of bacteria, that cause caries^{101, 146} and might have a carioprotective effect^{56, 136}. It was shown to have an anti-plaque effect in vitro and in-vivo (tests with volunteers)²³ It was also shown that Manuca honey, a very potent antimicrobial honey, has a positive effect against dental

plaque development and gingivitis and thus can be used in the place of refined sugar in the manufacture of candy¹⁰¹.

According to electron microscopic studies ingestion of honey does not cause erosion of tooth enamel as observed after drinking of fruit juice (pH 3.5). Ten minutes after consumption of fruit juice tooth erosion was seen, while 30 minutes after honey ingestion the erosion was only very weak. This effect can be explained only partially by the calcium, phosphorous and fluoride levels of honey, other colloidal honey components have to be also responsible⁶⁶.

Oral inflammation

Stomatitis is inflammation of the mouth mucose. Aphthous stomatitis, as well as as other oral lesions like recurrent herpes labialis, recurrent intraoral herpes, atrophic/erosive oral lichen planus, oral candidiasis and oral psoriasis can be successfully treated with honey. Honey significantly speeded resolution of the inflammatory and ulcerative oral lesions. It significantly lowered the pain sensation and reduced the duration of some lesions and increased the number of pain free days⁵⁸.

To treat the conditions listed above, smear the individual ulcers, boils or aphthae with honey - or simply trickle a good spoonful of honey in your mouth and then swirl it around the mouth, in order to reach as many sores as possible. (It will become runny, but honey does penetrate the tissues very quickly; and it seems it is precisely when diluted that its curative power is activated.) ¹

Plaqie and Gingivitis

A study investigated whether or not manuka honey with an antibacterial activity rated UMF 15 could be used to reduce dental plaque and clinical levels of gingivitis. A chewable "honey leather" was produced for this trial. Thirty volunteers were randomly allocated to chew or suck either the manuka honey product, or sugarless chewing gum, for 10 minutes, three times a day, after each meal. Plaque and gingival bleeding scores were recorded before and after the 21-day trial period. Analysis of the results indicated that there were statistically highly significant reductions in the mean plaque scores (0.99 reduced to 0.65; p=0.001), and the percentage of bleeding sites (48% reduced to 17%; p=0.001), in the manuka honey group, with no significant changes in the control group ⁶⁰

Against halitosis (malodour)

Manuka and acacia honey have been successfully used against halitosis (malodour) 138

Summarising the different findings, it can be concluded that honey is probably not as cariogenic as other sugars and in some cases can be also carioprotective, especially when strong antibacterial honey is ingested. However, for safety reasons, after consumption of honey it is advised to clean the teeth.

OTHER EXTERNAL APPLICATIONS BASED ON THE ANTIMICROBIAL ACTION OF HONEY

Besides the application in wounds and burns honey has also other external applications:

Against catheter infection of CKD

Honey can be used to fight Chronic Kidney Infections Disease (CKD), by application to the dialysis catheter This topic has been reviewed by Frances et al in 2015, in which it was reported that in three clinical studies application of honey to the could prevent successfully bacterial infections⁶¹ Topical medihoney can be successfully use against catheter infections⁷³

Against virus action on lips and genitals 14:

Apply honey on gauze auf critical point and change once a day

Against boils and furuncles

Mix liquid honey and flour 1:1, add a little water and brush with it affected area. Cover with gauze and leave it overnight.

Against muscle cramps

Cover affected area with honey, cover with gauze or cloth and fix it with adhesive plaster. Ev. cover with a warm wool cloth. Leave at least 2 hours.

Against bruises and contusions

Mix honey and olive oil 1:1 and cover with mixture affected area. Cover with gauze and leave for 4-6 hours.

Enhances post tonsilitis inflammation healing

Tualang honey from Malaysia enhances post tonsilitis inflammation healing process⁸⁵

Honey and cosmetics

A recent review summarises the used in honey in cosmetics. In cosmetic formulations, it exerts emollient, humectant, soothing, and hair conditioning effects, keeps the skin juvenile and retards wrinkle formation, regulates pH and prevents pathogen infections. Honey-based cosmetic products include lip ointments,

cleansing milks, hydrating creams, after sun, tonic lotions, shampoos, and conditioners. The used amounts range between 1 and 10%, but concentrations up to 70% can be reached by mixing with oils, gel, and emulsifiers, or polymer entrapment. Intermediate-moisture, dried, and chemically modified honeys are also used. Mechanisms of action on skin cells are deeply conditioned by the botanical sources and include antioxidant activity, the induction of cytokines and matrix metalloproteinase expression, as well as epithelial-mesenchymal transition in wounded epidermis⁴⁰

SIDE EFFECTS OF CANCER TREATMENTS

Most of the cancer research has been done in animal models (see chapter 7). The use of honey in clinical cancer treatments has been reviewed in 2008 by Bardy²⁷ and in 2009 by Orsolic¹¹⁴
The first reported use of honey in oncology patients was the topical application of 'household' honey to 12 patients with wound breakdown following radical excision of vulval carcinoma. Clearance of infection was observed within 3-6 days, and improved healing rates were recorded⁴¹. In a report from the Russian Academy of Medical Science, patients with uterine cancer undergoing radiotherapy and treated with 'honey laminolact' showed a significant decrease in the severity of radiation-induced intestinal morbidity¹⁴⁵

Honey treatment for prevention of oral mucositis

This topic has been reviewed in 2008 by Bardy. It has been pointed out that honey may be used for radiation-induced mucositis, radiotherapy-induced skin reactions, hand and foot skin reactions in chemotherapy patients and for oral cavity and external surgical wounds²⁷.

Bardy et al tested the effect of active manuka honey on radiation-induced mucositis. A total of 131 patients diagnosed with head and neck cancer who were having radiotherapy to the oral cavity or oropharyngeal area were recruited into the study, and were randomly allocated to take either manuka honey or placebo (golden syrup) 20 ml 4 times daily for 6 weeks. Mucositis was assessed according to the Radiation Therapy Oncology Group (RTOG) scale at baseline, weekly during radiotherapy, and twice weekly thereafter until the mucositis resolved. The patient's weight was recorded at the same time as the mucositis was assessed. Throat swabs to identify bacterial or fungal infections were taken at baseline, and during and after radiotherapy. There was no significant difference between honey and golden syrup in their effects on mucositis. Active manuka honey did not improve mucositis, but both the honey and the syrup seemed to be associated with a reduction in bacterial infections. Compliance was a problem after the onset of mucositis, which may have affected the findings²⁸.

Honey has a supportive effect on human patients who have undergone a cancer radiation therapy, decreasing radiation mucositis. Patients with head and neck cancer treated with radiation therapy were given honey. There was a significant reduction in the symptomatic grade 3/4 mucositis among honey-treated patients compared to controls; i.e. 20 versus 75%. The compliance of honey-treated group of patients was better than controls. Fifty-five percent of patients treated with topical honey showed no change or a positive gain in body weight compared to 25% in the control arm, the majority of whom lost weight ³¹.

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A randomized controlled clinical trial was conducted on 90 patients with acute lymphoblastic leukemia and oral mucositis grades 2 and 3. The mean age of enrolled patients was 6.9 years. The patients were assigned into 3 equal treatment groups: Honey, HOPE (honey, olive-oil propolis), and control groups. Recovery time in grade 2 mucositis was significantly reduced in the honey group as compared with either HOPE or controls (P < .05). In grade 3 mucositis, recovery time did not differ significantly between honey and HOPE (P = 0.61) but compared with controls, healing was faster with either honey or HOPE (P < .01). Generally, in both grades of mucositis, honey produced faster healing than either HOPE or controls (P < .05). Based on our results that showed that honey produced faster healing in patients with grade 2/3 chemotherapy-induced

mucositis, we recommend using honey and possibly other bee products and olive oil in future therapeutic trials targeting chemotherapy-induced mucositis³

Honey reduces chemoradiotherapy-induced mucositis in pediatric cancer patients⁷

Pediatric oncology

In paediatric oncology patients, the immune system is often suppressed by cytotoxic antineoplastic agents or radiation therapy and wound healing is impaired. In the Department of Paediatric Oncology at the Children's Hospital in the University of Bonn, MedihoneyTM has become a readily accepted treatment with a positive impact on patient and parent satisfaction¹⁴⁰.

Honey and chemotherapeutic drugs in combined supportive therapy

This use of honey has been reviewed⁸¹. Honey has been used to support chemotherapeutic action and reduce its side effects in myelosuppression, neutropenia etc.

Side effects of treatments of other cancers

Febrile neutropenia is a serious side effect of chemotherapy. Honey was administered to chemotherapy patients with neutropenia and was found that it reduced the need for colony-stimulating factors ¹⁶¹.

Therefore, the use of honey gauzes can be considered for the treatment of radiotherapy-induced dermatitis by radiotherapy of breast cancer patients¹⁰⁴

Topical application of honey can be used for the management of hhemotherapy induced oral stomatitis⁹⁷

The antitumor activity of honey can be explained by the antibacterial, antiinflammatory, immunodmodulating, anti-oxidant and probiotic effects of honey.

HONEY IN GASTROENTEROLOGY

According to the Muslim holy book "The Holy Hadith", dating back to the 8 th century AD the prophet Mohamed recommended honey against diarrhoea¹¹. Also, the Roman physician Celsus (ca. 25 AD) used honey as a cure for diarrhoea⁴². The use of honey for prevention and treatments of gastro-intestinal disorders such as peptic ulcers, gastritis, gastroenteritis has been reported in various books and publications from Eastern Europe^{80, 88, 95, 144} and from Arab countries¹³⁵ The applications of honey in traditional and modern medicine were reviewed in 2010¹

Ulcers and Gastritis

Honey is a potent inhibitor of the causing agent of peptic ulcers and gastritis, *Helicobacter pylori* ^{8, 21, 91, 116} In rats honey acted against experimentally induced gastric ulcers ^{18, 20, 65, 77}.

Honey is not involved on prostaglandin production, but has a stimulatory effect on the sensory nerves in the stomach that respond to capsaicin^{9, 16}. As a second mechanism of action has been postulated that this effect is due to the antioxidant properties of honey. Honey intake in rats prevented indomethacin-induced gastric lesions in rats by reducing the ulcer index, microvascular permeability and myeloperoxidase activity of the stomach¹⁰⁹. In addition, honey has been found to maintain the level of non-protein sulfhydryl compounds (e.g. glutathione) in gastric tissue subjected to factors inducing ulceration^{9, 16, 19}. A third mechanism of the gastroprotective effect of honey has been suggested by Beretta et al. It involves the salivary reduction of nitrate (NO3-) to nitrite (NO2-) and the intragastric formation of nitric oxide (NO), the latter involved in the preservation of the gastric mucosa capillaries and in boosting mucous production. Honeys contained NO2 and NO3, the concentration in honeydew honeys being higher than that of blossom honeys²⁹

Ingestion of dandelion honey was shown to reduce gastric juice acidity by 56% ²⁶. The gastric emptying of saccharides after ingestion of honey was slower than that of a mixture of glucose and fructose ¹²²

The effect of honey under clinical condition on more than 40 gastric ulcer patients was studied in a Russian hospital. Control treatments were with water. It was found that ingestion of 120 ml of 33 % honey solution by gastric ulcer patiens improves the micro capillary blood circulation, which can beneficially influence the gastric ulcers. Ingestion of 120 ml of 33 % honey warm honey solution decreases the acidity of the gastric juice, while the ingestion of the same amount and concentration of a cold honey solution increased the

acidity of the gastric juice. The sleep of the gastric ulcer patients was also improved by ingestion of 50 g honey before sleep. In order to decrease gastric juice acidity the author recommends the intake of warm honey solution 40 to 60 minutes before eating. The function of the gall bladder is improved by the ingestion of cool solution of 100 ml 50 % honey (13-15 $^{\circ}$ C) The author concludes that ingestion of warm honey ev. in combination with propolis ingestion, is a good way to treat gastric ulcers 54 .

A Bulgarian clinical study with dispeptic patients showed that honey consumption more than once weekly led to lower *Helicobacter pylori* rates of the patients³⁸

There are reports on healing of patients of suffering from gastritis, duodenitis and duodenal ulcers by intake of 30 ml of honey¹³⁵.

Clinical and animal studies have shown that honey reduces the secretion of gastric acid. Additionally, gastric ulcers have been successfully treated by the use of honey as a dietary supplement. An 80% recovery rate of 600 gastric ulcer patients treated with oral administration of honey has been reported. Radiological examination showed that ulcers disappeared in 59% of patients receiving honey⁷⁷.

Laxative effect against constipation

In certain cases, consumption of relatively large amounts of honey (50 to 100 g) can lead to a mild laxative effect in individual with insufficient absorption of honey fructose⁸³. Fructose is less readily absorbed in the intestinal tract than fructose together with glucose¹³¹. The mild laxative properties of honey are used for the treatment of constipation in Eastern Europe, China and the Near East. *However one should not give honey against constipation of infants younger than 1 year old because of the children botulism risk*.

Against acute gastroenteritis in children

A clinical study of honey treatment in infantile gastroenteritis was reported by Haffejee and Moosa. They found that by replacing the glucose (111 mmol/l) in the standard electrolyte-containing oral rehydration solution recommended by the World Health Organisation/UNICEF as well as the solution of electrolyte composition 48 mmol/l sodium, 28 mmol/l potassium, 76 mmol/l chloride ions, with 50 ml/l honey (29), the mean recovery times of patients (aged 8 days to 11 years) were significantly reduced. Honey was found to shorten the duration of diarrhoea in patients with bacterial gastroenteritis caused by organisms such as Salmonella, Shigella and E. coli. They recommended that honey was a safe substitute for glucose as long as it provided 111 mmol/l each of glucose and fructose. The high sugar content of honey means that it could be used to promote sodium and water absorption from the bowel⁶⁷.

In a later clinical trial in Egypt, honey added to oral rehydration solution promoted rehydration of the body and sped recovery from vomiting and diarrhea²

Against alcohol abuse

Animal experiments have shown that the administration of a honey solution via a tube in the stomach of rabbits prior to them being administered with 0.5 g ethanol per kg body weight, accelerated alcoholic oxidation. Honey administered subcutaneously or orally before oral administration of ethanol affords protection against gastric damage and reverses changes in pH induced by ethanol¹⁷

A controlled clinical trial demonstrated the use of fructose in the treatment of acute alcoholic intoxication. A small but significant increase occurred in the rate of fall of blood-ethanol levels and it was concluded that fructose may be beneficial in shortening the duration of alcoholic intoxication³⁹.

Positive effects of honey on ethanol intoxication such as disappearance in blood increase and of ethanol elimination rate has also been confirmed in studies with humans 112, 113.

Ingestion of both honey (2 g/kg body weight) and fructose, prevented the ethanol-induced transformation of erythrocytes of mice 160

Treatment of reflux oesophagus and heartburn

Honey has high density, high viscosity, and low surface tension, and therefore, can stay longer in the oesophagus as a coating on the mucus membrane, and can be used against reflux oesphagus ^{92, 141} and against heartburn ⁹²

Hepatitis and liver health

A positive effect of honey on hepatitis A patients was found after ingestion of clover and rape honey, causing a decrease of alanine aminotransferase activity (by 9 to 13 times) and of bilirubin production by 2.1 to 2.6 times²⁶.

Honey and digestion

Supplementation of honey in concentration of 2, 4, 6 and 8 g/100 g to protein fed to rats improved the protein and lipid digestibility ¹⁴².

The anti gastric ulcer and anti-gastritis effect of honey can be explained by its antibacterial and antiinflammatory action, as well as with its inhibitory effect on the acidity of the gastric juice. The positive effect of honey on nutrition function is also due to its prebiotic effect.

CARDIOVASCULAR HEALTH

The effects of ingestion of 75 g of natural honey by humans compared to the same amount of artificial honey (fructose plus glucose) or glucose on plasma glucose, plasma insulin, cholesterol, triglycerides (TG), blood lipids, C-reactive proteins and homocysteine, most of them being risk factors for cardiovascular diseases were studied in humans. Elevation of insulin and C-reactive protein was significantly higher after dextrose than after honey.

Dextrose reduced cholesterol and low-density lipoprotein-cholesterol (LDL-C). Artificial honey slightly decreased cholesterol and LDL-C and elevated TG. Honey reduced cholesterol, LDL-C, and TG and slightly elevated high-density lipoprotein-cholesterol (HDL-C). In patients with hyperlipidemia, artificial honey increased LDL-C, while honey decreased LDL-C¹³.

A similar study has been recently carried out in normal and overweight persons carrying a higher risk for a cardiovascular disease. These patients were given 70 g honey for 30 days. Results showed that honey caused a mild reduction in body weight (1.3%) and body fat (1.1%). Honey reduced total cholesterol (3%), LDL-C (5.8), triacylglycerole (11%), FBG (4.2%), and CRP (3.2%), and increased HDL-C (3.3%) in subjects with normal values, while in patients with elevated variables, honey caused reduction in total cholesterol by 3.3%, LDL-C by 4.3%, triacylglycerole by 19%, and CRP by 3.3% (p < 0.05). The conclusion of the authors is that consumption of natural honey reduces cardiovascular risk factors, particularly in subjects with elevated risk factors, and it does not increase body weight in overweight or obese subjects ¹⁵⁹. Honey decreases also platelet aggregation and blood coagulation ⁵

The above cited studies suggest small effects of honey on arteriosclerosis risk factors such as cholesterol, LDL-c and TG, the first studies being carried out with only 9 patients.

In a study with 30 persons and 30 controls it was shown that no significant decrease of cholesterol HDL and TG was encountered after ingestion of 75 g honey daily for a period of 14 days. While there were no effects in men, in women HDL values were increased in the controls having ingested sucrose, while in the honey group no increase was encountered, pointing out that honey has a positive effects in women ¹⁰⁸

The effect of honey intake on the blood risk factors was tested in diabetes 2 patients (controls with no intake). Body weight, total cholesterol, low-density lipoprotein-cholesterol and triglyceride decreased, while and high-density lipoprotein-cholesterol ratio increased significantly²⁵.

Honey can contain nitric oxide (NO) metabolites which are known cardiovascular disease risk indicators. Increased levels of nitric oxides in honey could have a protecting function in cardiovascular diseases. Total nitrite concentration in different biological fluids from humans, including saliva, plasma, and urine was measured after ingestion by humans of 80 g of honey. Salivary, plasma and urinary NO metabolites concentrations showed a tendency to increase ^{11, 13}. Different honey types contained various concentrations of NO metabolites, darker or fresh honeys containing more NO metabolites than light or stored honey. After heating, NO metabolites decreased in all the kinds of honey ^{11, 13}.

The cardiovascular effects of honey can be explained by its antioxidant and anti-inflammatory effects.

HONEY AGAINST COUGH

Small doses of honey, 1 to 2 tablespoons intake has been found to influence favourably cough^{50, 70, 155, 156} and also sleep¹⁵⁶ of children.

The dose of honey used was ½ tsp for 2-5 year olds, 1 teaspoon for the 6 to 11 year-olds and 2 tsp for 12 to 18 year-olds. Buckwheat honey was chosen in this study because of its high antioxidant properties. The same study shows that honey is more effective than a chemical anti cough syrup¹¹⁸

A review of the conducted clinical trials in the literature by Oduwole et al. in 2012 made the following conclusions: Honey may be better than 'no treatment' and diphenhydramine in the symptomatic relief of cough but not better than dextromethorphan. There is no strong evidence for or against the use of honey¹¹¹

A double-blind randomised controlled trial was conducted from 2008 to 2011 in Iran. Included in the study were 97 adults who had experienced persistent post-infectious cough (PPC) for more than three weeks. The participants were distributed into three groups. A jam-like paste was prepared which consisted of honey plus coffee for the first group ('HC'), prednisolone for the second group (steroid, 'S'), and guaifenesin for the third group (control, 'C'). The participants were told to dissolve a specified amount of their product in warm water and to drink the solution every eight hours for one week. Honey plus coffee was found to be the most effective treatment modality for PPC. The recipe for the honey-coffee mixture: 500 g honey mixed with 700 g instant coffee powder. The daily dose was 23.7 g of the mixture, taken 3 times a day¹²⁷.

Intake of 90 ml milk with 10 g honey was tested in Italy on children with non-specific acute cough, resulting in improvement, similar to the one achieved by standard anti-cough drugs⁹⁶

Honey relieved the cough in a clinical test with 568 children better than diphenhydramine but less than dextromethorphan⁷²

HONEY AND THE BRAIN

Al-Hymyari conducted a five-year pilot study involving 2290 cognitively intact subjects and 603 with mild cognitive impairment aged 65 and older. They were randomized to receive either one daily tablespoon of Middle East honey or placebo. They found that only 95 subjects who received honey compared to 394 who received placebo developed dementia. The study concluded that honey and its properties act as natural preventive therapies for both cognitive decline and dementia¹²

A study using Tualang honey was conducted on 102 healthy postmenopausal women randomly assigned to one of three groups; untreated control, estrogen plus progestin therapy and Tualang honey. The participants in Tualang honey and estrogen plus progestin therapy groups received 20 g Tualang honey supplement and Femoston conti 1/5 (1 mg 17- β estradiol and 5 mg dydrogesterone), respectively, daily for 16 weeks. The participants' memory and oxidative stress status were assessed pre- and post-intervention. Postmenopausal women who received Tualang honey showed improvement in their immediate memory but not in immediate memory after interference and delayed recall. This result was comparable with the memory improvement seen in women receiving estrogen plus progestin therapy¹¹⁷

The neurological effects of honey have been recently reviewed¹²⁸. The review points out that different honey components, mostly polyphenols, have neurological effects on the brain. There is also direct evidence: according to the original references cited in this review honey has following effects:

- 1. Honey is reported to be an important component of Brahma rasayan, an Ayurvedic formulation that is prescribed to extend the lifespan and improve memory, intellect, concentration, and physical strength
- 2. Postmenopausal women who received honey showed improvements in their immediate memory but not in immediate memory after interference or in delayed recall.

It has been claimed for a long time that honey influences beneficially human sleep, but there were no experiments to prove the claims. Ingestion of one to two table spoons of buckwheat honey (10-20 g) by children of 6 to 18 years (6-11 years old- one table spoon, 12-18 yeas old 2 table spoons) improved also the sleep of coughing children of 5 to 18 years (6-11 years old- one table spoon, 12-18 yeas old 2 table spoons) improved also the sleep of coughing children of 5 to 18 years (6-11 years old- one table spoon, 12-18 years old 2 table spoons) improved also the sleep of coughing children of 5 to 18 years (6-11 years old- one table spoon, 12-18 years old 2 table spoons) improved also the sleep of coughing children of 5 to 18 years (6-11 years old- one table spoon, 12-18 years old 2 table spoons) improved also the sleep of coughing children of 5 to 18 years (6-11 years old- one table spoon, 12-18 years old 2 table spoons) improved also the sleep of coughing children of 5 to 18 years (6-11 years old- one table spoon, 12-18 years old 2 table spoons) improved also the sleep of coughing children of 5 to 18 years (6-11 years old- one table spoon, 12-18 years old 2 table spoons) improved also the sleep of coughing children of 5 to 18 years (6-11 years old- one table spoon) improved also the sleep of coughing children of 5 to 18 years (6-11 years old- one table spoon) improved also the sleep of coughing children of 5 to 18 years (6-11 years old- one table spoon) improved also the sleep of 5 to 18 years (6-11 years old- one table spoon) improved also the sleep of 5 to 18 years (6-11 years old- one table spoon) improved also the sleep old- one table spoons of 5 to 18 years (6-11 years old- one table spoon) improved also the sleep old- one table spoons of 5 to 18 years (6-11 years old- one table spoons) improved also the sleep old- one table spoons of 5 to 18 years (6-11 years old- one table spoons) improved also table spoons of 5 to 18 years (6-11 years old- one table spoons) improved also table spoons old- one table spoo

According to a theoretical model for the influence of honey on sleep honey stabilizes blood sugar levels and contributes to the release of melatonin, the hormone required for recovery and rebuilding of body tissues during rest⁹³

HONEY AGAINST DIABETES?

A clinical trial in Egypt showed that that long-term consumption of honey might have positive effects on the metabolic derangements of type 1 diabetes⁴

OTHER HEALTH ENHANCING EFFECTS

Influenza and common cold

An Iranian study claims that intake of 50 g of honey daily reduces the length of the common cold by two days ¹²⁵.

The Ukranian doctors Frolov and Peresadin reported on a unique long term honey intake experiment. Frolov is the chair of the department of infectious diseases in the medical university of Luganska. All members of the department took 3 times a day, a total of 40-45 g of honey added to lukewarm tea. In the whole experiment 26 people took part in this unique experiment (n and number of years): n 5 for 20 y; n 6 for 15 y; n 8 for 10 y; n 5 for 5 to 10 y. During the whole experiment no other prophylactic was used. During the last 8 years of the experiment the department was in close contact with 40-60 patients with influenza and inflammation of the upper respiratory organs or with other infectious diseases like virus hepatitis, dysentery and even cholera. During the 20 year duration of the experiment no department member had any of the described diseases. In the immunological blood test it was found that the skin and the blood had an increased bactericidal activity, combined with very low microbial counts on the skin, while there were no pathogens in the whole area of the upper respiratory organs. And there was a control group to this experiment: a medical department, which was in close proximity of Frolov's test group, which had influenza or sore throat 3 to 4 times a year. This shows that a long term honey intake increases the anti-infectious immunity⁶².

Hay fever and allergic rhinitis

Another controversial possible application of honey is its use for preventing hay fever. Beekeepers claim that eating honey in the pre-vegetation season (i.e. during winter) will prevent or weaken hay fever symptoms.

A report by Croft presented evidence that daily ingestion during winter time of 10-20 g of honey resulted in improvements of hay fever symptoms in 16 out of 21 patients⁴⁷.

Münstedt and Kalder found a positive effect of honey ingestion on hay fever by means of questionnaire filled out by 29 beekeepers¹⁰⁷.

In another study it was shown that honey to which birch pollen were added was more effective of treating rhinitis against birch pollen than conventional honey 133

A 2002 clinical trial did not confirm the positive effects of honey ingestions, but honey was taken during the hay fever season and not before it 129.

Ingestion of 1 g/kg weight Tualang honey for 4 weeks by allergic rhinitis patients, complemented by an antihistamine drug treatment resulted in a significant improvement of allergic rhinitis of the honey supplemented group as compared to placebo (sugar plus antihistamine drug) treated controls ²⁴.

As hay fever is increasing in developed countries this issue should be faced with more clinical trials, carried out in a correct way. More research is necessary to clarify this possible effect of honey.

Infertility

In a preliminary announcement at the 2nd International Conference on the Medicinal Use of Honey in 2010 there is a preliminary announcement that intracervical injection of honey in women with chronic endocervitis was of positive therapeutic value both in terms of clinical cure and fertility enhancement¹. At

the same conference it was reported that honey has positive effect on the mechanical properties of the fetal membranes, may be through "collagen promoting action².

Anaemia

Remy Chauvin reviews different early works carrreid out by Theobald et al. and Frauenfelder and Errerich in Germany, Perez in Spain and Johnsen in Sutralia, carried out on 4-8 old infants. The dose given was one tea to one soup spoon in warm milk per day. The increase of blood haemoglobin was seen after one week of intake ⁴³

These clinical results are confirmed by experiments by Haydak et al. with rats, placed on a diet with milk and poor in iron. Only dark honeys, e.g. calluna, were capable of bringing blood haemoglobin values back to normal, while light honeys failed to do so⁶⁹

Menopause

An daily ingestion of table spoon of sunflower honey (about 20 g) for a period of 14 days by breast cancer patients improved the menopausal symptoms of 68 % of the patients, while a placebo effect of 25 % was expected 106

Longevity

A representative sub-sample of 665 men within the Caerphilly Cohort kept a weighed dietary record for seven days. Risk factors for vascular and other diseases in 41 men who recorded eating honey suggest that these men were on the whole healthier than the 624 men who had not recorded honey consumption. All-cause mortality during 25 years of follow-up was considerably lower in the men who had consumed honey, the hazard ratio, adjusted for a number of possible confounding factors, being 0.44 (95% confidence limits 0.23, 0.86; P<0.017). Because of the small number of subjects and of deaths in this study, further data from other large cohorts will be required before any effect upon mortality and other health effects of honey consumption can be adequately evaluated⁴⁵

Obesity

As a food rich in carbohydrates eating of large quantities of honey will of course cause obesity. However, eating of honey will cause less weight increase than the same quantities of sucrose^{32, 110, 159}

THE EXPERIENCE IN RUSSIA

Ludyansky, a chief doctor in a big Russian hospital, with life-long practice in apitherapy, has summarised the apitherapy knowledge in his monograph "Apitherapia" (in Russian)⁸⁸
Ludyanski summarises the medical uses of honey in his hospital in the following table:

Treated disease	Very good and good improvement	No improvement
Alopecia	11	5
Geriatry	59	-
Impotency	21	6
Inflammation of the vagina	21	5
Neurasthenia	60	-
Pediatrics	47	12
Prostatitis	24	5
Radicultis	47	15
Stomatology	43	16

HOW TO EXPLAIN THE USE OF HONEY IN MEDICINE

Therapeutic and health enhancing use	Biological rationale
Honey in healing of burns and wounds	Antibacterial, anti-inflammatory, antioxidant, osmotic and analgesic effects
Therapy of digestive diseases like peptic ulcers and gastritis	Antibacterial and anti-inflammatory effects
Against children diarrhoea	Antibacterial and anti-inflammatory effects
Improvement of gut microbial health and of digestion	Prebiotic effect
Improvement of immune reaction of the body	Immunoactivating effect
Regular intake improves cardiovascular health	Lowering of blood risk factors and specific heart conditions as extracystoles, arrhythmia and tachicardia
Long term ingestion of honey can reduce the risk of human cancer	Anticancerogenic effects
Positive glycemic nutritional effect. Can be used as a sweetener of people with diabetes type II and also probably type I	Some honeys have a low glycemic index: e.g. acacia honey. Other fructose rich honeys such as thyme, chestnut, heather and tupelo are good alternatives.
Use for the treatment of radiation-induced mucositis	Antibacterial and anti-inflammatory effects
Positive effect of honey ingestion on hepatitis A patients	Anti-inflammatory effect
Improvement of cough in children	Contact soothing effect, sweet substances, as a sweetener honey causes reflex salivation and increases airway secretions which may lubricate the airway and remove the trigger that causes a dry, nonproductive cough.

HONEY MASSAGE



Honey massage was developed in Tibet and Russia and is extensively described elsewhere 68, 151

Both liquid and crystalline honeys can be used.

1-2 tea spoons of liquid honey are applied on the back. Massagist puts hands puts hands onto this area and unglues the palms. Easy at first, "ungluing" the hands becomes more difficult with every move because the tension force increases. Massage lasts until the palms no longer stick to the massaged area, and the honey disappears from it. The actual duration depends on the type and quality of honey. Generally, honey massage lasts from 30 minutes.

IMPORTANCE OF THE HONEY TYPE



Due to different proportions of the possible sources, nectar and/or honeydew coming from a great variety of plants, no honey is completely the same as another one. This variability could be a handicap, given the market requirement for a consistent product, but when properly managed, it also could represent an opportunity for enhancing honey by offering to the consumer a number of typical products with special characteristics, according to the

particular botanical origin. Indeed, unifloral honeys are regarded as a more valuable class of honey, and botanical denominations are widely employed on the European market, often achieving higher prices than honey blends. Unifloral honeys have higher prices than blend honeys. In countries like France, Italy and Spain 30 to 50 % of the marketed honey is unifloral. In non-European countries, with the exception of the Manuka New Zealand honey, unifloral honeys have a smaller importance.

Information on European honeys is compiled in the special Apidologie Issue 35 from 2004. In Europe there are more than 100 plant species that can give origin to unifloral honey, most of them having only a local importance ¹¹⁹⁻¹²¹

Most biological and clinical studies reviewed above have been made with undetermined types of honeys and there are very few studies where comparisons have been done with different unifloral honeys. Here the fields will be reviewed where such studies have been carried out.

EVENTUAL HEALTH HAZARDS

See Chapter 8, Honey as nutrient and functional food.

HONEY INTAKE

The health enhancing effects in human adults, described in this report were mostly achieved after ingestion of 50 to 80 g of honey per day.

The health claims of honey which are reported below are valid for intakes of following amounts of honey:

- Adults: after ingestion of 50 to 80 g per day by adults,
- General (adults or infants): 0.8 g to 1.2 g honey per g human weight

The duration of honey ingestion for increase of physical performance and fitness

is very fast, and takes place <u>already 1 to 4 hours after intake</u>.

The health effects reported in the different publications reported above were measured mostly after $\underline{2}$ to $\underline{3}$ weeks of daily honey ingestion. Practical apitherapists suggest for health enhancing effects a daily honey ingestion for $\underline{1.5}$ to $\underline{2}$ months $\underline{^{65}, 89}$.

The normal daily allowance for carbohydrate sweeteners is 25 grams. Considering that the recommended amount of honey is quite high, intake of other sweeteners should be avoided. A normal intake of about 20-25 g per day will rather have a long term health enhancing effect.

References

- 1. ABDELLAH, F; ABDERRRAHIM, L (2010) Honey in gastrointestinal disorders, *In* Boukraa, L (ed.) *Honey in Traditional and Modern Medicine*, CRC Press Taylor and Francis Group; pp 160-186.
- 2. ABDULRHMAN, M A; MEKAWY, M A; AWADALLA, M M; MOHAMED, A H (2010) Bee Honey Added to the Oral Rehydration Solution in Treatment of Gastroenteritis in Infants and Children. *Journal of Medicinal Food* 13 (3): 605-609.
- 3. ABDULRHMAN, M; EL BARBARY, N S; AMIN, D A; EBRAHIM, R S (2012) Honey and a Mixture of Honey, Beeswax, and Olive Oil-Propolis Extract in Treatment of Chemotherapy-Induced Oral Mucositis: A Randomized Controlled Pilot Study. *Pediatric Hematology and Oncology* 29 (3): 285-292.
- 4. ABDULRHMAN, M M; EL-HEFNAWY, M H; ALY, R H; SHATLA, R H; MAMDOUH, R M; MAHMOUD, D M; MOHAMED, W S (2013) Metabolic Effects of Honey in Type 1 Diabetes Mellitus: A Randomized Crossover Pilot Study. *Journal of Medicinal Food* 16 (1): 66-72.
- 5. AHMED, A; KHAN, R A; AZIM, M K; SAEED, S A; MESAIK, M A; AHMED, S; IMRAN, I (2011) Effect of Natural Honey on Human Platelets and Blood Coagulation Proteins. *Pakistan Journal of Pharmaceutical Sciences* 24 (3): 389-397.
- 6. AHMED, A K; HOEKSTRA, M J; HAGE, J J; KARIM, R B (2003) Honey-medicated dressing: transformation of an ancient remedy into modern therapy. *Annals of Plastic Surgery* 50 (2): 143-147.
- 7. AL JAOUNI, S; HUSSEIN, A; AL MUHAYAWI, M; IBRAHIM, K; ELFIKI, I (2012) Honey reduces chemoradiotherapy-induced mucositis in pediatric cancer patients. *Critical Reviews in Oncology Hematology* 82: S17.
- 8. AL SOMAL, N; COLEY, K E; MOLAN, P C; HANCOCK, B M (1994) Susceptibility of *Helicobacter pylori* to the antibacterial activity of manuka honey. *Journal of the Royal Society of Medicine* 87 (1): 9-12.

- 9. AL SWAYEH, O A; ALI, A T (1998) Effect of ablation of capsaicin-sensitive neurons on gastric protection by honey and sucralfate. *Hepato Gastroenterology* 45 (19): 297-302.
- 10. AL WAILI, N S; SALOOM, K Y (1999) Effects of topical honey on post-operative wound infections due to gram positive and gram negative bacteria following caesarean sections and hysterectomies. *European journal of medical research* 4 (3): 126-130.
- 11. AL-BUKHAARI, M (1994) Holy Hadith (Sahih Al-Bukhari, Arabic). Kazi Publications Chicago (3rd. edition)
- 12. AL-HIMYARI, F A (2009) The use of honey as a natural preventive therapy of cognitive decline and dementia in the Middle East. *Alzheimers Dement* 5: 247.
- 13. AL-WAILI, N S (2004) Natural honey lowers plasma glucose, C-reactive protein, homocysteine, and blood lipids in healthy, diabetic, and hyperlipidemic subjects: Comparison with dextrose and sucrose. *Journal of Medicinal Food* 7 (1): 100-107.
- 14. AL-WAILI, N S (2004) Topical honey applications vs. acyclovir for the treatment of recurrent herpes simplex lesions. *Medical Science Monitor* 10 (8): 94-98.
- 15. AL-WAILI, N S; SALOM, K; AL-GHAMDI, A A (2011) Honey for Wound Healing, Ulcers, and Burns; Data Supporting Its Use in Clinical Practice. *ScientificWorldJournal*: 766-787.
- ALI, ATM (1995) Natural honey accelerates healing of indomethacin-induced antral ulcers in rats. Saudi Med.J. 16 (2): 161-166.
- 17. ALI, A T M M (1991) Prevention of ethanol-induced gastric lesions in rats by natural honey, and its possible mechanism of action. *Scandinavian Journal of Gastroenterology* 26: 281-288.
- 18. ALI, A T M M (1995) Natural honey exerts its protective effects against ethanol-induced gastric lesions in rats by preventing depletion of glandular nonprotein sulfhydryls. *Tropical Gastroenterol.* 16 (1): 18-26.
- 19. ALI, A T M M (1997) Natural honey prevents ischaemia-reperfusion-induced gastric mucosal lesions and increased vascular permeability in rats. *European Journal of Gastroenterology and Hepatology* 9 (11): 1101-1107.
- 20. ALI, A T M M (2003) Prevention of ammonia-induced gastric lesions in rats by natural honey. *Journal of Nutritional & Environmental Medicine* 13 (4): 239-246.
- 21. ALI, A T M M; CHOWDHURY, M N H; AL-HUMAYYD, M S (1991) Inhibitory effect of natural honey on Helicobacter pylori. *Trop.Gastroenterology* 12 (3): 139-143.
- 22. ALTMAN, N (2010) The honey prescription. Healing art press Rochester; Vermont 05767
- 23. APARNA, S; SRIRANGARAJAN, S; MALGI, V; SETLUR, K P; SHASHIDHAR, R; SETTY, S; THAKUR, S (2012) A Comparative Evaluation of the Antibacterial Efficacy of Honey In Vitro and Antiplaque Efficacy in a 4-Day Plaque Regrowth Model In Vivo: Preliminary Results. *Journal of Periodontology* 83 (9): 1116-1121.
- 24. ASHA'ARI, Z A; AHMAD, M Z; DIN, W S J W; HUSSIN, C M C; LEMAN, I (2013) Ingestion of honey improves the symptoms of allergic rhinitis: evidence from a randomized placebo-controlled trial in the East Coast of Peninsular Malaysia. *Annals of Saudi Medicine* 33 (5): 469-475.
- 25. BAHRAMI, M; ATAIE-JAFARI, A; HOSSEINI, S; FORUZANFAR, M H; RAHMANI, M; PAJOUHI, M (2009) Effects of natural honey consumption in diabetic patients: an 8-week randomized clinical trial. *International journal of food sciences and nutrition* 60 (7): 618-626.
- 26. BALTUSKEVICIUS, A; LAISKONIS, A; VYSNIAUSKIENE, D; CEKSTERYTE, V; RACYS, J (2001) Use of different kinds of honey for hepatitis A treatment and for reduction of increased acidity of gastric juice. *Zemdirbyste, Mokslo Darbai* 76: 173-180.
- 27. BARDY, J; SLEVIN, N J; MAIS, K L; MOLASSIOTIS, A (2008) A systematic review of honey uses and its potential value within oncology care. *Journal of Clinical Nursing* 17 (19): 2604-2623.

- 28. BARDY, J; MOLASSIOTIS, A; RYDER, W; MAIS, K; SYKES, A; YAP, B; LEE, L; KACZMARSKI, E; SLEVIN, N (2012) A double-blind, placebo-controlled, randomised trial of active manuka honey and standard oral care for radiation-induced oral mucositis. *British Journal of Oral & Maxillofacial Surgery* 50 (3): 221-226.
- 29. BERETTA, G; GELMINI, F; LODI, V; PIAZZALUNGA, A; FACINO, R M (2010) Profile of nitric oxide (NO) metabolites (nitrate, nitrite and N-nitroso groups) in honeys of different botanical origins: Nitrate accumulation as index of origin, quality and of therapeutic opportunities. *Journal of Pharmaceutical and Biomedical Analysis* 53 (3): 343-349.
- 30. BETTS, J A; MOLAN, P C (2001) A pilot trial of honey as a wound dressing has shown the importance of the way that honey is applied to wounds. *Paper of the European Wound Management Association Conference, Dublin, Eire*
- 31. BISWAL, B M; ZAKARIA, A; AHMAD, N M (2003) Topical application of honey in the management of radiation mucositis. A preliminary study. *Supportive Care in Cancer* 11 (4): 242-248.
- 32. BISWAS, B K (2009) Effects of Honey on Feed Consumption and Body Weight of Sprague-Dawley and Obese Rats. *Journal of the American Association for Laboratory Animal Science* 48 (5): 613.
- 33. BOGDANOV, S; GALLMANN, P; STANGACIU, S; CHERBULIEZ, T (2006) Bienenprodukte und Gesundheit. *AlpForum* 41: 3-50.
- 34. BOUKRAA, L E (2010) Honey in Traditional and Modern Medicine. CRC Press Taylor and Francis Group
- 35. BOUKRAA, L (2010) Honey in burn and wound management, *In* Boukraa, L (ed.) *Honey in Traditional and Modern Medicine*, CRC Press Taylor and Francis Group; pp 125-153.
- 36. BOUKRAA, L; SULAIMAN, S A (2010) Honey Use in Burn Management: Potentials and Limitations. *Forschende Komplementarmedizin* 17 (2): 74-80.
- 37. BOWEN, W H; LAWRENCE, R A (2005) Comparison of the cariogenicity of cola, honey, cow milk, human milk, and sucrose. *Pediatrics* 116 (4): 921-926.
- 38. BOYANOVA, L; ILIEVA; J.GERGOVA, G; R.MITOV I (2015) Honey and green/black tea consumption may reduce the risk of Helicobacter pylori infection. *Diagnostic Microbiology and Infectious Disease* 82: 85-86.
- 39. BROWN, S; FOREST, J; ROSCOE, P (1972) A controlled trial of fructose in the treatment of acute alcoholic intoxication. *Lancet* 2: 898-890.
- 40. BURLANDO, B; CORNARA, L (2013) Honey in dermatology and skin care: a review. *Journal of Cosmetic Dermatology* 12 (4): 306-313.
- 41. CAVANAGH, D; BEAZLEY, J; OSTAPOWICZ, F (1970) Radical operation for carcinoma of the vulva. A new approach to wound healing. *Journal of Obstetrics and Gynaecology* 77 (11): 1037-1040.
- 42. CELSUS, C (1935) De medicina. Heinemann London, UK
- 43. CHAUVIN, R (1968) Action physiologique et therapeutique des produits de la ruche *Traite de biologie de l'abeille*, Masson; Paris; pp 116-154.
- 44. COHEN, H A; ROZEN, J; KRISTAL, H; LAKS, Y; BERKOVITCH, M; UZIEL, Y; KOZER, E; POMERANZ, A; EFRAT, H (2012) Effect of honey on nocturnal cough and sleep quality: A double-blind, randomized, placebo-controlled study. *Pediatrics*: 1-8.
- 45. COOPER, R; FEHILY, A M; PICKERING, J; ERUSALIMSKY, J; ELWOOD, P (2010) Honey, Health and Longevity. *Current Aging Science*, 3: 239-241.
- 46. COOPER, R A; HALAS, E; MOLAN, P C (2002) The efficacy of honey in inhibiting strains of Pseudomonas aeruginosa from infected burns. *Journal of Burn Care and Rehabilitation* 23 (6): 366-370.

- 47. CROFT, L (1990) Honey and hay fever: a report on the treatment of hay fever with honey.
- 48. CUTTING K.; WHITE R. (2002) Maceration of the skin: 1. The nature and causes of skin maceration. *Journal of Wound Care* 11: 275-278.
- 49. CUTTING, K F (2007) Honey and contemporary wound care: An overview. *Ostomy/Wound Management* 53 (11): 49-54.
- 50. DARBY-STEWART, A; DACHS, R; GRABER, M A (2009) Honey as a Treatment for Cough in Children. *American Family Physician* 80 (2): 120-121.
- 51. DAVIES, P (2005) Recent Clinical Usage of Honey in the Treatment of Wounds A Review. *Wounds* 5 (1): 14-22.
- 52. DECAIX, C (1976) Comparative study of sucrose and honey. *Le Chirurgien-dentiste de France* 46 (285-286): 59-60.
- 53. DESCOTTES, B (2009) Cicatrisation par le miel, l'éxpérience de 25 années. Phytotherapie 7: 112-116.
- 54. DUBTSOVA, E (2009) Clinical studies with bee products for therapy of some nutritional diseases (in Russian). Central Moscow Institute of Gastroenterology Moscow; pp 1-38.
- 55. DUNFORD, C; COOPER, R; MOLAN, P (2000) Using honey as a dressing for infected skin lesions. *NT Plus* 96 (14): 7-9.
- 56. EDGAR, W M; JENKINS, G N (1974) Solubility-reducing agents in honey and partly-refined crystalline sugar. *British Dental Journal* 136: 7-14.
- 57. EFEM, S E E (1988) Clinical observations on the wound healing properties of honey 183. *British Journal of Surgery* 75: 679-681.
- 58. EL-HADDAD, S; SHAWAF, M (2013) Effect of honey for treatment of some common oral lesions: Follow up of 50 cases. *Journal of Dentistry and Oral Hygiene* 5: 55-61.
- 59. EMARAH, M H (1982) A clinical study of the topical use of bee honey in the treatment of some occular diseases. *Bulletin of Islamic Medicine* 2 (5): 422-425.
- 60. ENGLISH, H K; PACK, A R; MOLAN, P C (2004) The effects of manuka honey on plaque and gingivitis: a pilot study. *Journal of the International Academy of Periodontology* 6 (2): 63-67.
- 61. FRANCIS, A; CHO, Y; JOHNSON, D (2015) Honey in the Prevention and Treatment of Infection in the CKD Population: A Narrative Review. *eCam* http://dx.doi.org/10.1155/2015/261425
- 62. FROLOV, V M; PERESSADIN, N A (2006) Honey against influenza and sore throat. *Pcelovodstvo* 10 (529): 52-53.
- 63. GANNABATHULA, S; SKINNER, M A; ROSENDALE, D; GREENWOOD, J M; MUTUKUMIRA, A N; STEINHORN, G; STEPHENS, J; KRISSANSEN, G W; SCHLOTHAUER, R C (2012) Arabinogalactan proteins contribute to the immunostimulatory properties of New Zealand honeys. *Immunopharmacology and Immunotoxicology* 34 (4): 598-607.
- 64. GEORGE, N M; CUTTING, K F (2007) Antibacterial honey (Medihoney (TM)): in-vitro activity against clinical isolates of MRSA, VRE, and other multiresistant gram-negative organisms including Pseudomonas aeruginosa (vol 19, pg 231, 2007). *Wounds-A Compendium of Clinical Research and Practice* 19 (10): A10.
- 65. GHARZOULI, K; AMIRA, S; GHARZOULI, A; KHENNOUF, S (2002) Gastroprotective effects of honey and glucose-fructose-sucrose-maltose mixture against ethanol-, indomethacin-, and acidified aspirin-induced lesions in the rat. *Experimental and toxicologic pathology* 54 (3): 217-221.

- 66. GROBLER, S R; DU TOIT, I J; BASSON, N J (1994) The effect of honey on human tooth enamel in vitro observed by electron microscopy and microhardness measurements. *Archives of Oral Biology* 39: 147-153.
- 67. HAFFEJEE, I E; MOOSA, A (1985) Honey in the treatment of infantile gastroenteritis. *British Medical Journal* 290: 1866-1867.
- 68. HARNISCH, G (2008) Die Entgiftungsmassage mit Honig. Altes russisches Heilwissen neu entdeckt leicht anzuwenden. Lorber U. Turm Verlag; 88 pp
- 69. HAYDAK, M H; PALMER, L S; TANQUARY, M C (1942) The role of honey in the prevention and cure of nutritional anemia in rats. *Journal of Pediatrics* 21 (6): 763-768.
- 70. HEPPERMANN, B; JONES, J S (2009) Honey for the Symptomatic Relief of Cough in Children with Upper Respiratory Tract Infections. *Emergency Medicine Journal* 26 (7): 522-523.
- 71. IRISH, J; CARTER, D; BLAIR, S (2005) Honey kills some of our most dangerous microbial enemies *Apimondia abstracts Ireland 2005*, Dublin; pp 124.
- 72. JARRETT, B; CORBO, J; JASON M. (2015) Not so sweet: Honey for acute cough in children. *Evidence Based Practice* https://hdl.handle.net/10355/45978
- 73. JOHNSON D.W.; VAN EPS C.; MUDGE DW.; WIGGINS KJ; ARMSTRONG K; HAWLEY CM; CAMPBELL SB; ISBEL NM; NIMMO GR; GIBBS H (2005) Randomized, controlled trial of topical exit-site application of honey (Medihoney) versus mupirocin for the prevention of catheter-associated infections in hemodialysis patients. *J.Am.Soc.Nephrol.* 16: 1456-1462.
- 74. JONES, R (2001) Honey and healing through the ages, *In* Munn, P; Jones, R (eds) *Honey and healing*, IBRA International Bee Research Association; Cardiff, GB; pp 1-4.
- 75. JULL, A B; RODGERS, A; WALKER, N (2008) Honey as a topical treatment for wounds. *Cochrane Database of Systematic Reviews* (Issue 4. Art. No.: CD005083. DOI: 10.1002/14651858.CD005083.pub2.)
- 76. JULL, A B; WALKER, N; DESHPANDE, S (2013) Honey as a topical treatment for wounds. *Cochrane Database of Systematic Reviews* (2)
- 77. KANDIL, A; EL-BANBY, M; ABDEL-WAHED, K; ABDEL-GAWWAD, M; FAYEZ, M (1987) Curative properties of true floral and false nonfloral honeys and induced gastric ulcers. *J.Drug.Res.Egypt* 17 (1-2): 103-106.
- 78. KARAYIL, S; DESHPANDE, S D; KOPPIKAR, G V (1998) Effect of honey on multidrug resistant organisms and its synergistic action with three common antibiotics. *Journal of Postgraduate Medicine* 44 (4): 93-96.
- 79. KHISMATULLINA, N (2005) Apitherapy. Perm, Russia
- 80. KHOTKINA, M L (1955) Honey as part of therapy for patients with stomach ulcers. *Collection of papers from the Irkutsk State Medical Institute*: 252-262.
- 81. KRISHNA, R (2005) Therapeutic uses of Honey in Ayurveda.: http://www.ezilon.com/articles/articles/356-1/Therapeutic-uses-of-Honey-in-Ayurveda.
- 82. KUMAR, A; SHARMA, V K; SINGH, H P; PRAKASH, P; SINGH, S P (1993) Efficacy of some indigenous drugs in tissue repair in buffaloes. *Indian Veterinary Journal* 70 (1): 42-44.
- 83. LADAS, S D; RAPTIS, S A (1999) Honey, fructose absorption, and the laxative effect. *Nutrition* 15 (7-8): 591-592.
- 84. LAHANAS, M (2010) Examples of Ancient Greek Medical Knowledge.: accessed 8.2.2010 on http://www.mlahanas.de/Greeks/Med.htm.

- 85. LAZIM, N M; ABDULLAH, B; SALIM, R (2013) The effect of Tualang honey in enhancing post tonsillectomy healing process. An open labelled prospective clinical trial. *International Journal of Pediatric Otorhinolaryngology* 77 (4): 457-461.
- 86. LEE, D S; SINNO, S; KHACHEMOUNE, A (2011) Honey and Wound Healing An Overview. *American Journal of Clinical Dermatology* 12 (3): 181-190.
- 87. LETCHUMANAN, P; RAJAGOPALAN, R; KAMARUDDIN, M Y (2013) Posttonsillectomy pain relief and epithelialization with honey. *Turkish Journal of Medical Sciences* 43 (5): 851-857.
- 88. LUDYANSKII, E A (1994) *Apitherapy* 1231. Poligrafist Vologda, Russia
- 89. MADDOCKS, S E; JENKINS, R E; ROWLANDS, R S; PURDY, K J; COOPER, R A (2013) Manuka honey inhibits adhesion and invasion of medically important wound bacteria in vitro. *Future Microbiology* 8 (12): 1523-1536.
- 90. MAJTAN, J; BOHOVA, J; GARCIA-VILLALBA, R; TOMAS-BARBERAN, F A; MADAKOVA, Z; MAJTAN, T; MAJTAN, V; KLAUDINY, J (2013) Fir honeydew honey flavonoids inhibit TNF-alpha-induced MMP-9 expression in human keratinocytes: a new action of honey in wound healing. *Archives of Dermatological Research* 305 (7): 619-627.
- 91. MANYI-LOH, C E; CLARKE, A M; MUNZHELELE, T; GREEN, E; MKWETSHANA, N F; NDIP, R N (2010) Selected South African Honeys and Their Extracts Possess In Vitro Anti-Helicobacter pylori Activity. *Archives of Medical Research* 41 (5): 324-331.
- 92. MATH, M V; KHADKIKAR, R M; KATTIMANI, Y R (2013) Honey A nutrient with medicinal property in reflux oesophagitis. *Indian Journal of Medical Research* 138: 1020-1021.
- 93. MCINNIS, M (2008) The Uniqueness of Honey - its impact on Human Metabolism and its role in Restorative Sleep, *First International Symposium on honey and health*, Sacremento
- 94. MCLOONEA, P; WARNOCKB, M; FYFEB, L (2015) Honey: A realistic antimicrobial for disorders of the skin. *Journal of Microbiology, Immunology and Infection* doi:10.1016/j.jmii.2015.01.009
- 95. MENSHIKOV, F K; FEIDMAN, S I (1949) Curing stomach ulcers with honey. *Sovetskaya Meditsing* 10: 13-14.
- 96. MICELI SOPOA, S; GRECOA, M; MONACOA, S; VARRASI, G; DI LORENZO, G; SIMEONE, G (2015) Effect of multiple honey doses on non-specific acute cough in children. An open randomised study and literature review. *Allergologia et Immunopathologia* 43: 449-455.
- 97. MOHAMED, S A; SHEBL, A; WEHEIDA, S M (2012) The Effect of Topical Application of Honey on Management of Chemotherapy Induced Oral Stomatitis. *Life Science Journal-Acta Zhengzhou University Overseas Edition* 9 (4): 5128-5134.
- 98. MOLAN, P (2002) Not all honeys are the same for wound healing. Eur Tissue Repair Soc Bulletin 9: 5-6.
- 99. MOLAN, P (2005) Mode of action, *In* White, R; Molan, P; Copper, R (eds) *Honey: A modern wound management product*, Wounds UK; Aberdeen; pp 1-23.
- 100. MOLAN, P C (2001) Honey as a topical antibacterial agent for treatment of infected wounds. http: www.World Wide Wounds.com.Report: 1-13.
- 101. MOLAN, P C (2001) Honey for oral health. Journal of Dental Research 80 (special issue): 1-130.
- 102. MOLAN, P C (2001) Why honey is effective as a medicine 1. Its use in modern medicine, *In* Munn, P; Jones, R (eds); pp 5-13. (82. edition)
- 103. MOLAN, P C (2002) Re-introducing honey in the management of wounds and ulcers theory and practice. Ostomy/Wound Management 48 (11): 28-40.

- 104. MOOLENAAR, M; POORTER, R L; VAN DER TOORN, P P; LENDERINK, A W (2006) The effect of honey compared to conventional treatment on healing of radiotherapy--induced skin toxicity in breast cancer patients. *Acta Oncologica* 45: 623-624.
- 105. MOORE, O A; SMITH, L A; CAMPBELL, F; SEERS, K; MCQUAY, H J; MOORE, R A (2001) Systematic review of the use of honey as a wound dressing. *BMC Complementary and Alternative Medicine* 1 (1): 2.
- 106. MUENSTEDT, K; VOSS, B; KULMER, U; SCHNEIDER U; HÜBNER, U (2015) Bee pollen and honey for the alleviation of hot flushes and other menopausal symptoms in breast cancer patients. *Mol Clin Oncol* DOI: 10.3892/mco.2015.559
- 107. MÜNSTEDT, K; KALDER, M (2010) Honey as a treatment option for rhinoconjunctivitis. JAAS 2: 145-148.
- 108. MÜNSTEDT, K; HOFFMANN, S; HAUENSCHILD, A; BULTE, M; VON GEORGI, R; HACKETHAL, A (2009) Effect of Honey on Serum Cholesterol and Lipid Values. *Journal of Medicinal Food* 12 (3): 624-628.
- 109. NASUTI, C; GABBIANELLI, R; FALCIONI, G; CANTALAMESSA, F (2006) Antioxidative and gastroprotective activities of anti-inflammatory formulations derived from chestnut honey in rats. *NUTRITION RESEARCH* 26 (3): 130-137.
- 110. NEMOSECK, T M; CARMODY, E G; FURCHNER-EVANSON, A; GLEASON, M; LI, A; POTTER, H; REZENDE, L M; LANE, K J; KERN, M (2011) Honey promotes lower weight gain, adiposity, and triglycerides than sucrose in rats. *NUTRITION RESEARCH* 31 (1): 55-60.
- 111. ODUWOLE, O; MEREMIKWU, M M; OYO-ITA, A; UDOH, E E (2012) Honey for acute cough in children. *Cochrane Database of Systematic Reviews* (3)
- 112. ONYESOM, I (2004) Effect of Nigerian citrus (Citrus sinensis Osbeck) honey on ethanol metabolism. *Samj South African Medical Journal* 94 (12): 984-986.
- 113. ONYESOM, I (2005) Honey-induced stimulation of blood ethanol elimination and its influence on serum triacylglycerol and blood pressure in man. *Annals of Nutrition and Metabolism* 49 (5): 319-324.
- 114. ORSOLIC, N (2009) Honey and Cancer. JAAS 1 (4): 93-103.
- 115. ORYAN, A; ZAKER, S R (1998) Effects of topical application of honey on cutaneous wound healing in rabbits. *Journal of veterinary medicine.A, Physiology, pathology, clinical medicine* 45 (3): 181-188.
- 116. OSATO, M S; REDDY, S G; GRAHAM, D Y (1999) Osmotic effect of honey on growth and viability of Helicobacter pylori. *Digestive diseases and sciences* 44 (3): 462-464.
- 117. OTHMAN, Z; SHAFIN, N; ZAKARIA, R; HUSSAIN, N H N; MOHAMMAD, W M Z W (2011) Improvement in immediate memory after 16 weeks of tualang honey. *Menopause* 18: 1219-1224.
- 118. PAUL, I M; BEILER, J; MCMONAGLE, A; SHAFFER, M L; DUDA, L; BERLIN, C M (2007) Effect of honey, dextromethorphan, and no treatment on nocturnal cough and sleep quality for coughing children and their parents. *Archives of Pediatrics & Adolescent Medicine* 161 (12): 1140-1146.
- 119. PERSANO ODDO, L; PIANA, L; BOGDANOV, S; BENTABOL, A; GOTSIU, P; KERKVLIET, J; MARTIN, P; MORLOT, M; VALBUENA, A O; RUOFF, K; VON DER OHE, K (2004) Botanical species giving unifloral honey in Europe. *Apidologie* 35 (special issue): 82-93.
- 120. PERSANO ODDO, L; PIRO, R (2004) Main European unifloral honeys: descriptive sheets. *Apidologie* 35 (special issue): S38-S81.
- 121. PIAZZA, M G; PERSANO ODDO, L (2004) Bibliographical review of the main European unifloral honeys. *Apidologie* 35 (special issue): S94-S111.

- 122. POKORN, D; VUKMIROVIC, V (1978) Velocity of gastric emptying of saccharides after administering honey, apicompleks and pure invert sugar, *IIIe Symposium International d'Apitherapie*, 11-15 Septembre 1978, Portoroz, Yougoslavie, Apimondia, Bukarest, 1978: pp 277-279.
- 123. POSTMES, T; VANDEPUTTE, J (1999) Recombinant growth factors or honey? Burns 25 (7): 676-678.
- 124. POTSCHINKOVA, P (1999) Apitherapie: Die Heilkraft von Honig and Co. Ehrenwirth Verlag München
- 125. POURAHMAD, M; SOBHANIAN, S (2009) Effect of Honey on the Common Cold. *Arch Med Res* 40: 224-225.
- 126. PRAKASH, S; RATNA, R (2014) Honey in Ayurvedic Medicine, *In* Boukraa, L (ed.) *Honey in Traditional and Modern Medicine*, CRC Press Taylor and Francis Group; pp 13-20.
- 127. RAEESSI, M A; ASLANI, J; RAEESSI, N; GHARAIE, H; ZARCHI, A A K; RAEESSI, F (2013) Honey plus coffee versus systemic steroid in the treatment of persistent post-infectious cough: a randomised controlled trial. *Primary Care Respiratory Journal* 22 (3): 325-330.
- 128. RAHMAN, M M; GAN, S H; KHALIL, M I (2014) Neurological Effects of Honey: Current and Future Prospects. *Evidence-based complementary and alternative medicine*
- 129. RAJAN, T V; TENNEN, H; LINDQUIST, R L; COHEN, L; CLIVE, J (2002) Effect of Ingestion of Honey on Symptoms of Rhinoconjunctivitis. *Annals of allergy, asthma & immunology* 88 (2): 198-203.
- 130. RAYNAUD, A; GHEZALI, L; GLOAGUEN, V; LIAGRE, B; QUERO, F; PETIT, J M (2013) Honey-induced macrophage stimulation: AP-1 and NF-kappa B activation and cytokine production are unrelated to LPS content of honey. *International immunopharmacology* 17 (3): 874-879.
- 131. RIBY, J E; FUJISAWA, T; KRETCHMER, N (1993) Fructose absorption. *The American Journal of Clinical Nutrition* 58 (5): 748-753.
- 132. RIPPON, M; JONES, D (2005) A Review of the Physical Performance Characteristics of Honey-based Wound Dressings and Ointments. *Wounds, Honey Supplement* 5: 50-60.
- 133. SAARINEN, K; JANTUNEN, J; HAAHTELA, T (2011) Birch Pollen Honey for Birch Pollen Allergy A Randomized Controlled Pilot Study. *International Archives of Allergy and Immunology* 155 (2): 160-166.
- 134. SALEHI, A; JABARZARE, S; NEURMOHAMADI, M; KHEIRI, S; RAFIEIAN-KOPAEI M (2014) A Double Blind Clinical Trial on the Efficacy of Honey Drop in Vernal Keratoconjunctivitis. *eCam* http://dx.doi.org/10.1155/2014/287540
- 135. SALEM, S N (1981) Honey regimen in gastrointestinal disorders. Bulletin of Islamic Medicine 1: 358-362.
- 136. SELA, M O; SHAPIRA, L; GRIZIM, I; LEWINSTEIN, I; STEINBERG, D; GEDALIA, I; GROBLER, S R (1998) Effects of honey consumption on enamel microhardness in normal versus xerostomic patients. *Journal of Oral Rehabilitation* 25 (8): 630-634.
- 137. SHANNON, I L; EDMONDS, E J; MADSEN, K O (1979) Honey: Sugar content and cariogenicity. *Journal of dentistry for children*: 29-33.
- 138. SHIGA, H; JO, A; TERAO, K; NAKANO, M; OSHIMA, T; MAEDA, N (2010) Decrease of halitosis by intake of manuka honey, *General Session of IADR Barcelona*, 14.July2010
- 139. SIEDENTOPP, W (2009) Honey: Effective Against Inflammation, Cough and Hoarseness. *Deutsche Zeitschrift fuer Akkupunktur* 52: DOI: 10.1016/j.dza.2009.10.004.
- 140. SIMON, A; SOFKA, K; WISZNIEWSKY, G; BLASER, G; BODE, U; FLEISCHHACK, G (2006) Wound care with antibacterial honey (Medihoney) in pediatric hematology-oncology. *Supportive Care in Cancer* 14 (1): 91-97.

- 141. SINGH, P; SINGH, N; SENGUPTA, S; PALIT, G (2013) Honey A nutrient with medicinal property in reflux oesophagitis Response. *Indian Journal of Medical Research* 138: 1021.
- 142. SIRNIK, V; KOCH, V; GOLOB, T (1978) The influence of honey on the digestibility of nutritive substances for albin rats (L'influence du miel sur la digestibilité des substances nutritives chez le rat albinos), *IIIe Symposium International d'Apitherapie, 11-15 Septembre 1978, Portoroz, Yougoslavie*, Apimondia, Bukarest, 1978: pp 286-290.
- 143. SIU-WAN, IP (2007) Honey in Chinese Culture. Malays J Med Sci 14: 101-127.
- 144. SLOBODIANIUK, A A; SLOBODIANIUK, M S (1969) Complex treatment of gastritis patients with high stomach secretion in combination with (and without) a 15-20% solution of honey, *Resorts of Bashkiria in the Service of Health.*, Ufa, Bashkir. Khniz. izd.-vo: pp 249-253.
- 145. SMIRNOVA, I I; FILATOVA, E I; SUVOROV, A N; BYLINSKAIA, E N (2000) [The use of therapeutic/prophylactic dragee "honey laminolact" in radiotherapy of uterine tumors]. *Voprosy Onkologii* 46 (6): 748-750.
- 146. STEINBERG, D; KAINE, G; GEDALIA, I (1996) Antibacterial effect of propolis and honey on oral bacteria. *American Journal of Dentistry* 9 (6): 236-239.
- 147. STEPHEN-HAYNES, J (2004) Evaluation of honey impregnated tulle dressing in primary care. *Brit J Community Nurs, Wound Care Supplement*: S21-S27.
- 148. SUBRAHMANYAM, M (1997) A prospective randomised clinical and histological study of superficial burn wound healing with honey and silver sulfadiazine. *Burns* 24: 157-161.
- 149. SUBRAHMANYAM, M; SAHAPURE, A; NAGANE, N (2001) Effects of topical application of honey on burn wound healing. *Ann Burns Fire Disasters* 14: 143-145.
- 150. SUBRAHMANYAM, M; SAHAPURE, A; NAGANE, N (2003) Free radical control the main mechanism of the action of honey in burns. *Ann Burns Fire Disasters* 16: 135-138.
- 151. TSCHENZE, V (2001) Russisch-Tibetische Honigmassage.; 92 pp (Books on Demand, Videel. edition)
- 152. UWAYDAT, S; JHA, P; TYTARENKO, R; BROWN, H; WIGGINS, M; BORA, P S; BORA, N S (2011) The Use of Topical Honey in the Treatment of Corneal Abrasions and Endotoxin-Induced Keratitis in an Animal Model. *Current eye research* 36 (9): 787-796.
- 153. VAN DEN BERG AJ; VAN DEN WORM E; VAN UFFORD HC; HALKES SB; HOEKSTRA MJ; BEUKELMAN CJ (2008) An in vitro examination of the antioxidant and anti-inflammatory properties of buckwheat honey. *J Wound Care* 17: 172-174-176-178.
- 154. WAHDAN, H A (1998) Causes of the antimicrobial activity of honey. Infection 26 (1): 26-31.
- 155. WARREN, M D; COOPER, W O (2008) Honey improves cough in children compared to no treatment. *Journal of Pediatrics* 152 (5): 739-740.
- 156. WARREN, M D; PONT, S J; BARKIN, S L; CALLAHAN, S T; CAPLES, T L; CARROLL, K N; PLEMMONS, G S; SWAN, R R; COOPER, W O (2007) The effect of honey on nocturnal cough and sleep quality for children and their parents 226. Archives of Pediatrics & Adolescent Medicine 161 (12): 1149-1153.
- 157. WHITE, R; MOLAN, P; COPPER, R; EDS. (2005) *Honey: A modern wound management product*. Wounds UK Aberdeen; 160 pp
- 158. WOOD, B; RADEMAKER, M; MOLAN, P (1997) Manuka honey, a low cost leg ulcer dressing. *The New Zealand medical journal* 110 (1040): 107.
- 159. YAGHOOBI, N; AL WAILI, N; GHAYOUR-MOBARHAN, M; PARIZADEH, S M R; ABASALTI, Z; YAGHOOBI, Z; YAGHOOBI, F; ESMAEILI, H; KAZEMI-BAJESTANI, S M R; AGHASIZADEH, R; SALOOM, K Y; FERNS, G A A (2008) Natural honey and cardiovascular risk factors; Effects on

Book of Honey, Chapter 9

- blood glucose, cholesterol, triacylglycerole, CRP, and body weight compared with sucrose. *Thescientificworldjournal* 8: 463-469.
- 160. YAMADA, S; ITOH, E; MURAKAMI, Y; ASANO, M (1999) Prevention of ethanol-induced erythrocyte transformations by fructose and natural honey in low alcohol tolerance mice. *Pathophysiology* 6: 163-170.
- 161. ZIDAN, J; SHETVER, L; GERSHUNY, A; ABZAH, A; TAMAM, S; STEIN, M; FRIEDMAN, E (2006) Prevention of chemotherapy-induced neutropenia by special honey intake. *Medical Oncology* 23 (4): 549-552.
- 162. ZUMLA, A; LULAT, A (1989) Honey a remedy rediscovered. *Journal of the Royal Society of Medicine* 82: 384-385.